FILE 'REGISTRY' ENTERED AT 16:55:19 ON 05 MAR 2003 L8 0 S EC 1.11.1.12/CN L9 1 S PHOSPHOLIPID HYDROPEROXIDE GLUTATHIONE PEROXIDASE/CN FILE 'CA' ENTERED AT 16:57:00 ON 05 MAR 2003 1098 S L1 L10 231 S L9 L11 S 97089-70-8/REG# FILE 'REGISTRY' ENTERED AT 16:57:29 ON 05 MAR 2003 1 S 97089-70-8/RN L12 FILE 'CA' ENTERED AT 16:57:30 ON 05 MAR 2003 231 S L12 L13 L14 18 S L13 AND INHIBITOR L15 18 S L13 AND L14

> s 19 L11 231 L9

=> s 97089-70-8

## REG1stRY INITIATED

Substance data SEARCH and crossover from CAS REGISTRY in progress... Use DISPLAY HITSTR (or FHITSTR) to directly view retrieved structures.

L13 231 L12

=> s 113 and inhibitor

396059 INHIBITOR

420358 INHIBITORS

644275 INHIBITOR

(INHIBITOR OR INHIBITORS)

L14 18 L13 AND INHIBITOR

=> s 113 and 114

L15 18 L13 AND L14

=> d ti 1-18

- L15 ANSWER 1 OF 18 CA COPYRIGHT 2003 ACS
- TI Novel aspects related to biosynthesis and biological actions of hepoxilins: interrelationship with phospholipid hydroperoxide glutathione peroxidase (PHGPx)
- L15 ANSWER 2 OF 18 CA COPYRIGHT 2003 ACS
- TI Involvement of reactive oxygen species in arsenite-induced downregulation of phospholipid hydroperoxide glutathione peroxidase in human epidermoid carcinoma A431 cells
- L15 ANSWER 3 OF 18 CA COPYRIGHT 2003 ACS
- TI Effects of signaling molecules, protein phosphatase inhibitors and blast pathogen (Magnaporthe grisea) on the mRNA level of a rice (Oryza sativa L.) phospholipid hydroperoxide glutathione peroxidase (OsPHGPX) gene in seedling leaves
- L15 ANSWER 4 OF 18 CA COPYRIGHT 2003 ACS
- TI Gene expression profiles associated with osteoblast differentiation
- L15 ANSWER 5 OF 18 CA COPYRIGHT 2003 ACS
- TI Saccharomyces cerevisiae expresses three phospholipid hydroperoxide glutathione peroxidases
- L15 ANSWER 6 OF 18 CA COPYRIGHT 2003 ACS
- TI Phospholipid hydroperoxide glutathione peroxidase protects against singlet oxygen-induced cell damage of photodynamic therapy
- L15 ANSWER 7 OF 18 CA COPYRIGHT 2003 ACS
- TI Criteria for the identification of housekeeping genes and their use as internal standards in the measurement of levels of gene expression
- L15 ANSWER 8 OF 18 CA COPYRIGHT 2003 ACS
- TI Method for identifying toxic agents in liver tissues using differential gene expression
- L15 ANSWER 9 OF 18 CA COPYRIGHT 2003 ACS

- L9 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS
- RN 97089-70-8 REGISTRY
- CN Peroxidase, glutathione (phospholipid hydroperoxide-reducing) (9CI) (CA INDEX NAME)

## OTHER NAMES:

- CN E.C. 1.11.1.12
- CN Phospholipid hydroperoxide glutathione peroxidase
- CN Selenoperoxidase
- MF Unspecified
- CI MAN
- LC STN Files: AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CAPLUS, CASREACT, EMBASE, TOXCENTER, USPATFULL

## \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

- 231 REFERENCES IN FILE CA (1962 TO DATE)
  - 4 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 231 REFERENCES IN FILE CAPLUS (1962 TO DATE)

- TI Involvement of phospholipid hydroperoxide glutathione peroxidase in the modulation of prostaglandin D2 synthesis
- L15 ANSWER 10 OF 18 CA COPYRIGHT 2003 ACS
- TI Method to search for male antifertility drugs based on PHGPx activity determination
- L15 ANSWER 11 OF 18 CA COPYRIGHT 2003 ACS
- TI Identification of a lipoxygenase inhibitor in A431 cells as a phospholipid hydroperoxide glutathione peroxidase
- L15 ANSWER 12 OF 18 CA COPYRIGHT 2003 ACS
- Overexpression of phospholipid hydroperoxide glutathione peroxidase suppressed cell death due to oxidative damage in rat basophile leukemia cells (RBL-2H3)
- L15 ANSWER 13 OF 18 CA COPYRIGHT 2003 ACS
- TI Purification of a cytosolic enzyme from human liver with phospholipid hydroperoxide glutathione peroxidase activity
- L15 ANSWER 14 OF 18 CA COPYRIGHT 2003 ACS
- TI Superoxide dismutase gene mutations as causes of neurodegenerative diseases and compounds and methods for the diagnosis, treatment, and prevention of the diseases
- L15 ANSWER 15 OF 18 CA COPYRIGHT 2003 ACS
- TI Selenoperoxidase-mediated cytoprotection against merocyanine 540-sensitized photoperoxidation and photokilling of leukemia cells
- L15 ANSWER 16 OF 18 CA COPYRIGHT 2003 ACS
- TI Antioxidant effect of Ebselen (PZ 51): peroxidase mimetic activity on phospholipid and cholesterol hydroperoxides vs free radical scavenger activity
- L15 ANSWER 17 OF 18 CA COPYRIGHT 2003 ACS
- TI Lethal damage to murine L1210 cells by exogenous lipid hydroperoxides: protective role of glutathione-dependent selenoperoxidases
- L15 ANSWER 18 OF 18 CA COPYRIGHT 2003 ACS
- TI Different effects of Triton X-100, deoxycholate, and fatty acids on the kinetics of glutathione peroxidase and phospholipid hydroperoxide glutathione peroxidase
- => d bib ab 10 13 14 15 18
- L15 ANSWER 10 OF 18 CA COPYRIGHT 2003 ACS
- AN 133:189863 CA
- TI Method to search for male antifertility drugs based on PHGPx activity determination
- IN Flohe, Leopold; Ursini, Fulvio
- PA Germany
- SO PCT Int. Appl., 33 pp. CODEN: PIXXD2
- DT Patent
- LA English
- FAN.CNT 1
  - PATENT NO. KIND DATE APPLICATION NO. DATE
- PI WO 2000053800 A1 20000914 WO 2000-EP1878 20000306
  - W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HU, ID, IL, IS, JP, KE, KG, KP,
    - KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO,
    - NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA,

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UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
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1159445 A1 20011205 EP 2000-910774 20000306 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO

PRAI EP 1999-103960 A 19990309 WO 2000-EP1878 W 20000306

AB The invention relates to a method to search for male antifertility drugs based on activity detn. of phospholipid hydroperoxide glutathione peroxidase (PHGPx) derived from human tissue or human cells or from related mammalian species.

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L15 ANSWER 13 OF 18 CA COPYRIGHT 2003 ACS
- AN 122:26500 CA
- TI Purification of a cytosolic enzyme from human liver with phospholipid hydroperoxide glutathione peroxidase activity
- AU Chambers, Stephen J.; Lambert, Nigel; Williamson, Gary
- CS Food Molecular Biochemistry Department, Institute Food Research, Norwich, NR4 7UA, UK
- SO International Journal of Biochemistry (1994), 26(10/11), 1279-86 CODEN: IJBOBV; IŜSN: 0020-711X
- PB Elsevier
- DT Journal
- LA English
- Phospholipid hydroperoxide glutathione peroxidase (PHGPx) is a AB selenoprotein which inhibits peroxidn. of microsomes. The human enzyme, which may play an important role in protecting the cell from oxidative damage, has not been purified or characterized. PHGPx was isolated from human liver using ammonium sulfate fractionation, affinity chromatog. on bromosulfophthalein-glutathione-agarose, gel filtration on Sephadex G-50, anion exchange chromatog. on Mono Q resin and high resoln. gel filtration on Superdex 75. The protein was purified about 112,000-fold, and 12 .mu.g was obtained from 140 g of human liver with a 9% yield. PHGPx was active on hydrogen peroxide, cumene hydroperoxide, linoleic acid hydroperoxide and phosphatidylcholine hydroperoxide. The mol. wt., as estd. from non-denaturing gel filtration, was 16,100. The turnover no. (37.degree., pH 7.6) on (.beta.-(13-hydroperoxy-cis-9, trans-11-octadecadienoyl)-.gamma.-palmitoyl)-L-.alpha.-phosphatidylcholine was 91 mol mol-1 s-1. reported for pig PHGPx, the activity of the enzyme from human liver on cumene hydroperoxide and on linoleic acid hydroperoxide was inhibited by deoxycholate. In the presence of glutathione, the enzyme was a potent inhibitor of ascorbate/Fe induced lipid peroxidn. in microsomes derived from human B lymphoblastic AHH-1 TK .+-. CHol cells but not from human liver microsomes. Human cell line microsomes contained no detectable PHGPx activity. However, microsomes prepd. from human liver contained 0.009 U/mg of endogenous PHGPx activity, which is 4-5 times the activity required for max. inhibition of lipid peroxidn. when pure PHGPx was added back to human lymphoblastic cell microsomes. PHGPx from human liver exhibits similar properties to previously described enzymes with PHGPx activity isolated from pig and rat tissues, but does not inhibit peroxidn. of human liver microsomes owing to a high level of PHGPx activity already present in these microsomes.
- L15 ANSWER 14 OF 18 CA COPYRIGHT 2003 ACS
- AN 121:246338 CA
- TI Superoxide dismutase gene mutations as causes of neurodegenerative diseases and compounds and methods for the diagnosis, treatment, and prevention of the diseases
- IN Brown, Robert; Horvitz, H. Robert; Rosen, Daniel R.
- PA General Hospital Corp., USA; Massachusetts Institute of Technology

SO PCT Int. Appl., 98 pp.

CODEN: PIXXD2

DT Patent LA English

FAN.CNT 1

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APPLICATION NO. DATE
               KIND DATE
    PATENT NO.
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   WO 9419493
                 A1 19940901
                                 WO 1994-US2089 19940228
PΙ
       W: CA, JP
       RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
                 A 19981201
                                 US 1993-23980
   US 5843641
                                               19930226
    CA 2157041
                     19940901
                 AA
                                  CA 1994-2157041 19940228
                 A1 19951213
                                 EP 1994-910183 19940228
    EP 686203
       R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE
    JP 08510377 T2 19961105 JP 1994-519309 19940228
                                  US 1995-486953 19950607
   US 5849290
                 Α
                     19981215
PRAI US 1993-23980
                     19930226
   US 1994-204052
                     19940228
   WO 1994-US2089
                      19940228
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AB Disclosed is the family of genes responsible for the neurodegenerative diseases, particularly amyotrophic lateral sclerosis (ALS). Methods and compds. for the diagnosis, prevention, and therapy of the disease are also disclosed. Uses of the compds. in the prepn. of diagnostic and therapeutic medicaments are also provided. Fourteen different SOD1 missense mutations in 16 different familial ALS families were identified. The mutations were identified by PCR followed by single-strand conformational polymorphism anal. The most common single mutation was an Ala-4 to Val substitution in exon 1. This mutation reduced the total SOD activity by 50% compared to normal controls. Addnl. polymorphisms were identified in exons 3 and 4 as well as in intron 3. Some of these mutations are detectable by restriction fragment length polymorphism.

- L15 ANSWER 15 OF 18 CA COPYRIGHT 2003 ACS
- AN 117:229272 CA
- TI Selenoperoxidase-mediated cytoprotection against merocyanine 540-sensitized photoperoxidation and photokilling of leukemia cells
- AU Lin, Fubao; Geiger, Peter G.; Girotti, Albert W.
- CS Dep. Biochem., Med. Coll. Wisconsin, Milwaukee, WI, 53226, USA
- SO Cancer Research (1992), 52(19), 5282-90 CODEN: CNREA8; ISSN: 0008-5472
- DT Journal
- LA English
- AB Photodynamic therapy with the lipophilic sensitizing dye merocyanine 540 (MC540) is a promising new approach for extracorporeal purging of neoplastic cells from autologous remission bone marrow grafts. Resistance-conferring cellular defenses against the cytotoxic effects of MC540/photodynamic therapy have not been well characterized. This study focuses on the cytoprotective effects of the glutathione-dependent selenoperoxidases GPX and PHGPX, which can detoxify a wide variety of hydroperoxides, including lipid-derived species (LOOHs). Murine leukemia L1210 cells were grown in 1% serum media without [L.cntdot.Se(-)] and with [L.cntdot.Se(+)] selenium supplementation. L.cntdot.Se(-) cells expressed 10-20-fold lower GPX and PHGPX activities than L.cntdot.Se(+) controls and were markedly more sensitive to MC540-mediated photoperoxidn. (LOOH formation) and clonally assessed photokilling. Susceptibility of L.cntdot.Se(-) cells to photoperoxidn. and photokilling could be fully reversed to L.cntdot.Se(+) levels by replenishing Se, and partially reversed by treating with Ebselen, a selenoperoxidase mimetic. Altered lipid compn., greater uptake of MC540, and defective catabolism of H2O2 were all ruled out as possible factors in the elevated photosensitivity of L.cntdot.Se(-) cells. Human leukemia K562 cells (capable of expressing PHGPX but not GPX) exhibited 5-10-fold lower PHGPX activity under Se-deficient relative to Se-sufficient conditions. Although MC540 uptake (nmol/mg lipid) by K562 and L1210 cells was essentially the same, the

former were more resistant to photoinactivation. However, like murine counterparts, Se-deficient cells were more susceptible to photoperoxidn. and photokilling than Se-sufficient controls. These results clearly demonstrate that GPX and/or PHGPX in L1210 cells and PHGPX in K562 cells play an important cytoprotective role during photooxidative stress. Whether membrane damage due to lipid photoperoxidn. is causally related to cell death is not certain; however, the parallel effects of Se deficiency on LOOH formation and cell killing are at least consistent with this possibility.

- L15 ANSWER 18 OF 18 CA COPYRIGHT 2003 ACS
- AN 106:63476 CA
- TI Different effects of Triton X-100, deoxycholate, and fatty acids on the kinetics of glutathione peroxidase and phospholipid hydroperoxide glutathione peroxidase
- AU Maiorino, Matilde; Roveri, Antonella; Gregolin, Carlo; Ursini, Fulvio
- CS Inst. Biol. Chem., Univ. Padova, Padua, 35131, Italy
- SO Archives of Biochemistry and Biophysics (1986), 251(2), 600-5 CODEN: ABBIA4; ISSN: 0003-9861
- DT Journal
- LA English
- The effects of Triton X 100, deoxycholate, and fatty acids were studied on AB the 2 steps of the ping-pong reaction catalyzed by Se-dependent glutathione peroxidases. The study was carried out by analyzing the single progression curves where the specific glutathione oxidn. was monitored by using glutathione reductase and NADPH. Although the classical glutathion peroxidase was inhibited only by Triton, the newly discovered phospholipid hydroperoxide glutathione peroxidase (from pig heart) was inhibited by deoxycholate and by unsatd. fatty acids. The kinetic anal. showed that in the case of glutathione peroxidase only the interaction of the lipophilic peroxidic substrate was hampered by Triton, indicating that the enzyme is not active at the interface. Phospholipid hydroperoxide glutathione peroxidase activity measured with linoleic acid hydroperoxide as substrate on the other hand, was not stimulated by Triton concns. which were shown to stimulate the activity with phospholipid hydroperoxides. Furthermore a slight inhibition was apparent at high Triton concns., and the effect could be attributed to a surface diln. of the substrate. Deoxycholate and unsatd. fatty acids were not inhibitory to glutathione peroxidase but inhibited both steps of the peroxidic reaction of phospholipid hydroperoxide glutathion-peroxidase, in the presence of either amphiphilic or hydrophilic substrates. This inhibition pattern suggests an interaction of anionic detergents with the active site of this enzyme. These results are in agreement with the different roles played by these peroxidases in the control of lipid peroxide concns. in the cells. Whereas glutathione peroxidase reduces the peroxides in the water phase (mainly H2O2), the new peroxidase reduces the amphiphilic peroxides, possibly at the water-lipid interface.

## => d his

(FILE 'HOME' ENTERED AT 16:12:55 ON 05 MAR 2003)

FILE 'CA' ENTERED AT 16:13:16 ON 05 MAR 2003

- L1 1098 S APYRASE
- L2 47648 S ALKALINE PHOSPHATASE
- L3 5762 S ADENOSINE DEAMINASE
- L4 7 S L1 AND L2 AND L3

FILE 'WPIDS' ENTERED AT 16:21:49 ON 05 MAR 2003

E SUGIYAMA ATSUSHI/AU 25

- L5 154 S E1 OR E2
- L6 34 S APYRASE
- L7 2 S L6 AND L5